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- (54) **THREE-DIMENSIONAL BIOLOGICAL SCAFFOLD COMPROMISING POLYMER WAVEGUIDES**
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(57) **ABSTRACT**

A three-dimensional biological scaffold. The scaffold includes at least three sets of polymer waveguides extending along at least three respective directions. The at least three sets of polymer waveguides interpenetrate each other at a plurality of nodes to form a self-supporting structure. In some embodiments, the polymer waveguides may be bio-degradable. In still some embodiments, the three-dimensional biological scaffold may include one or more coating layers for covering surfaces of the polymer waveguides.

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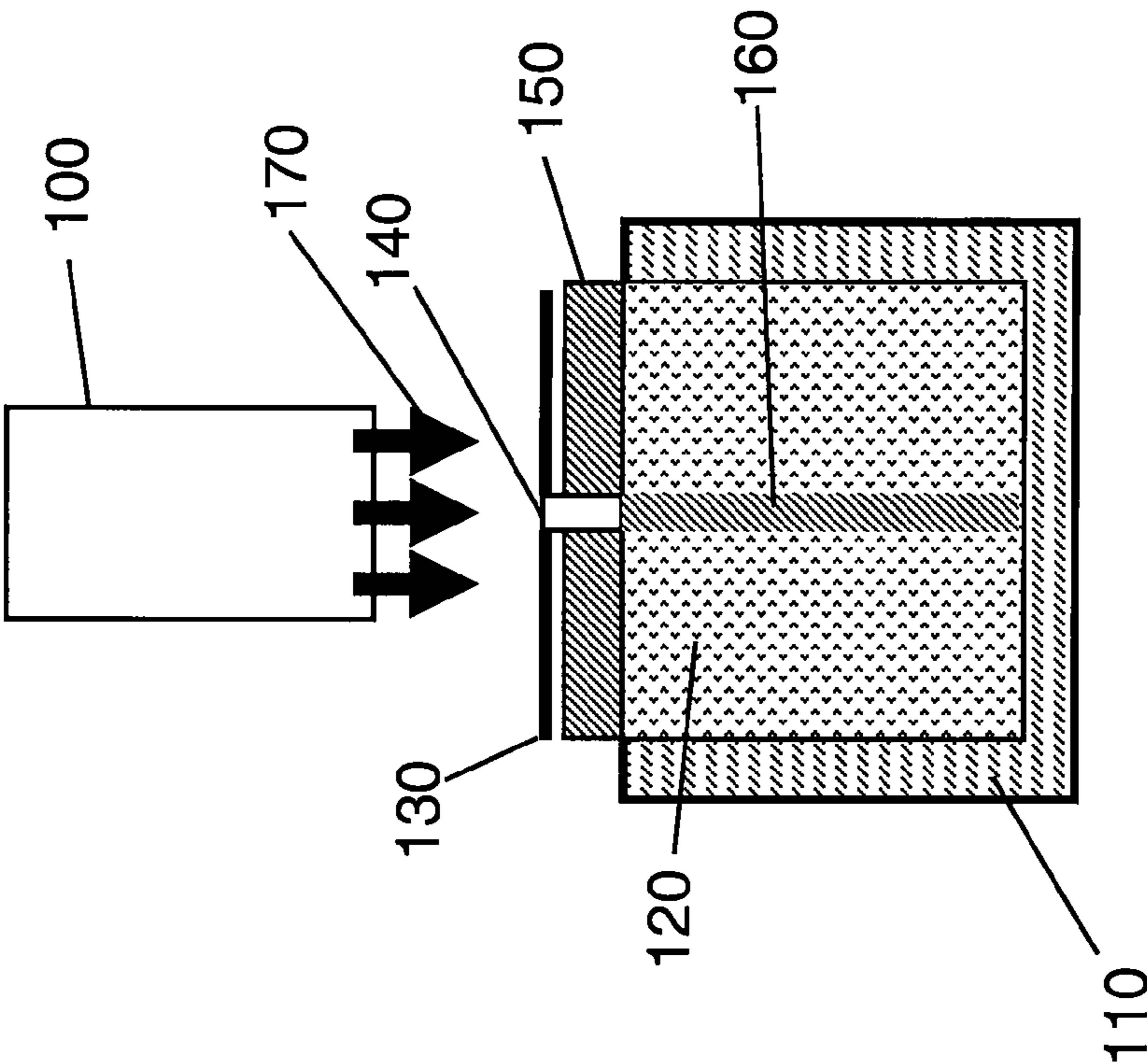


FIG. 1

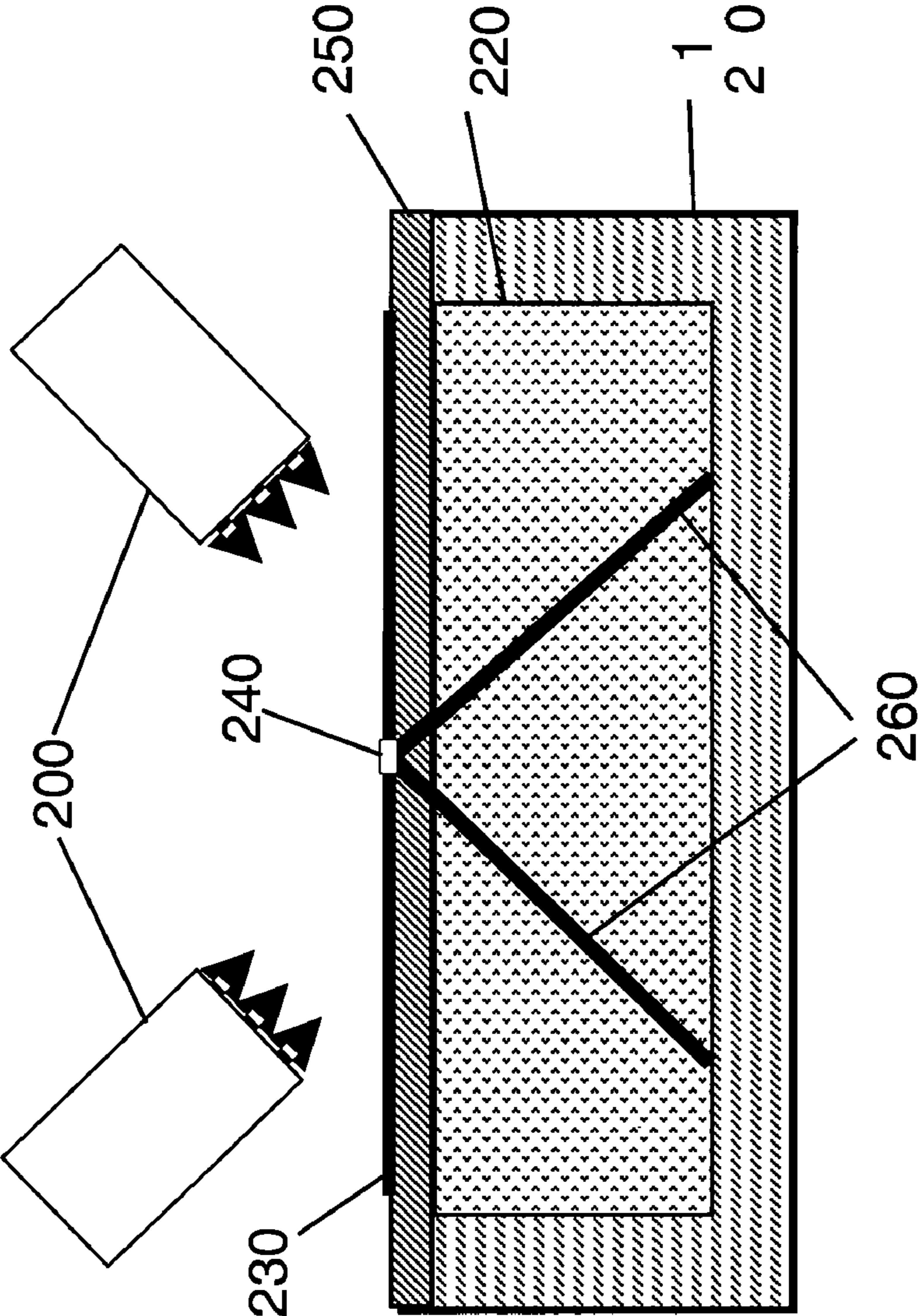


FIG. 2

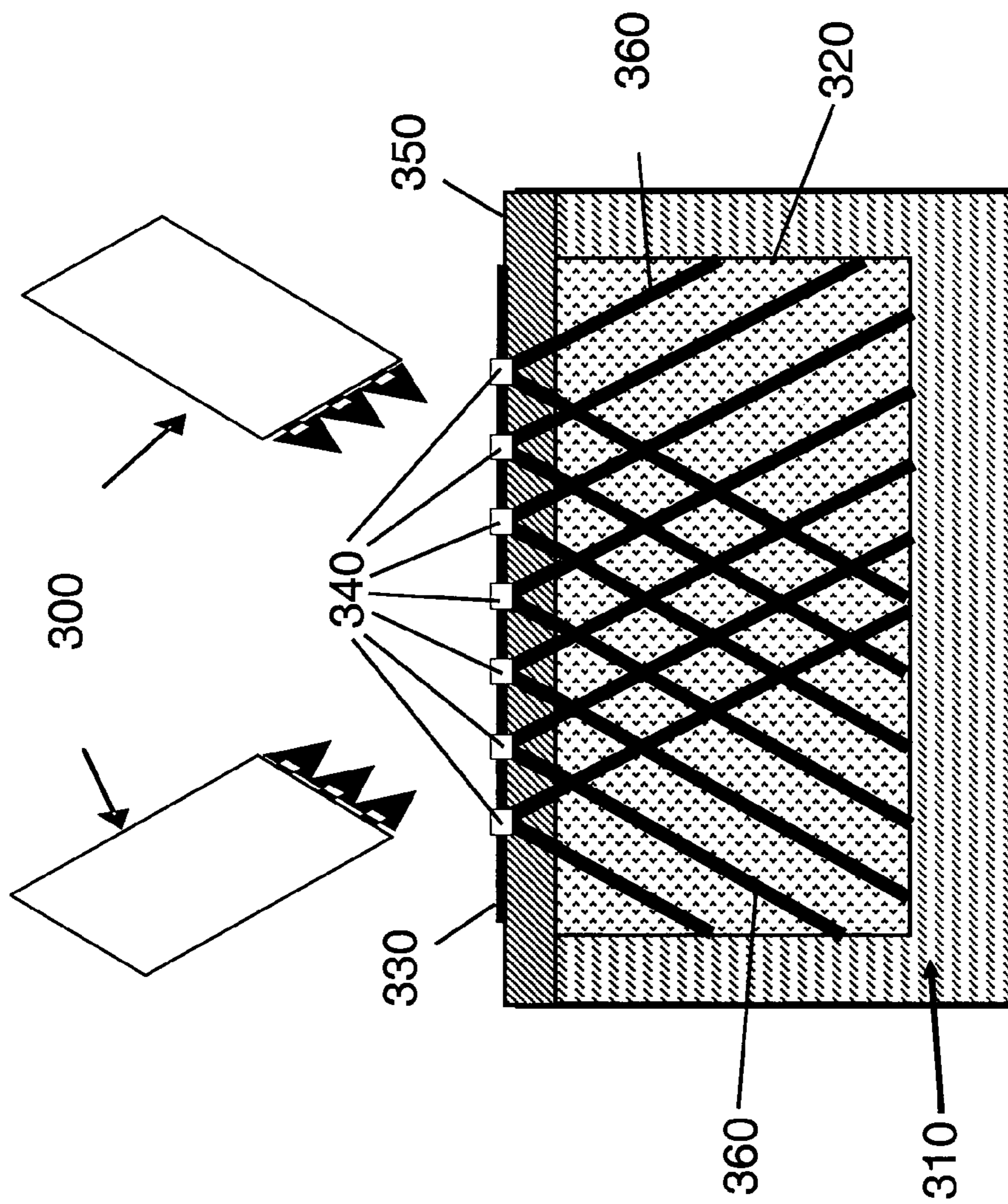


FIG. 3

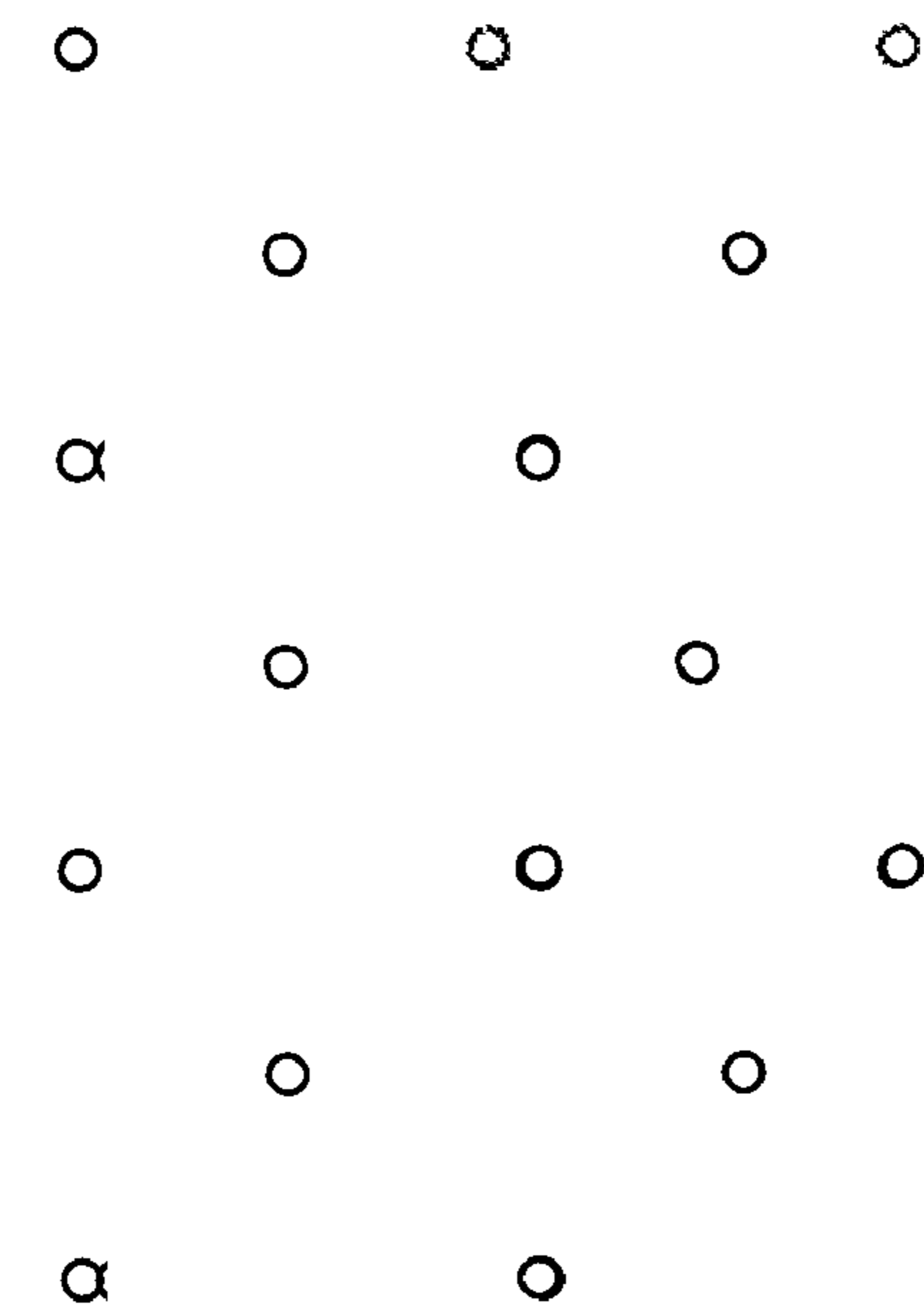
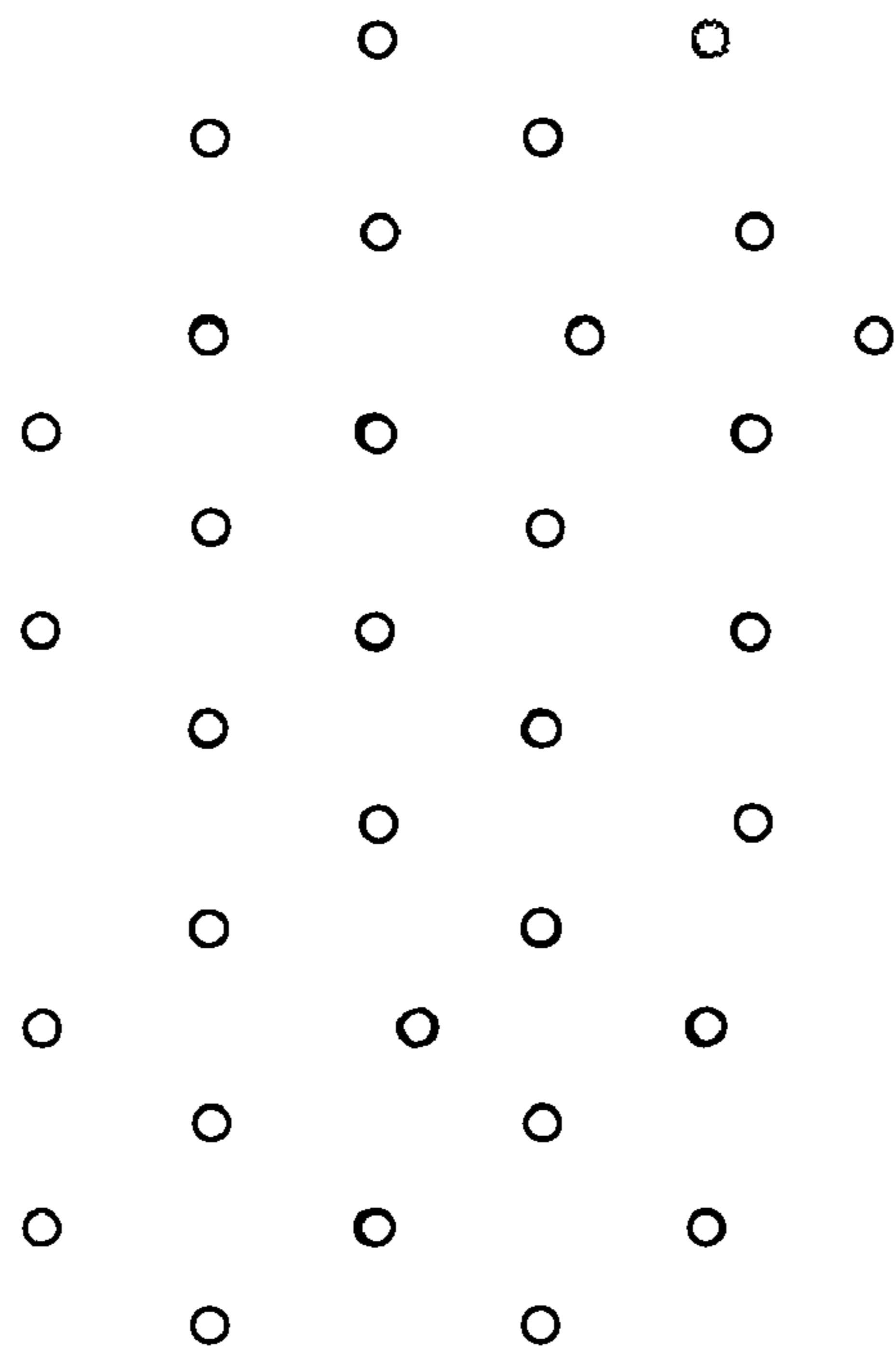


FIG. 4b

FIG. 4a

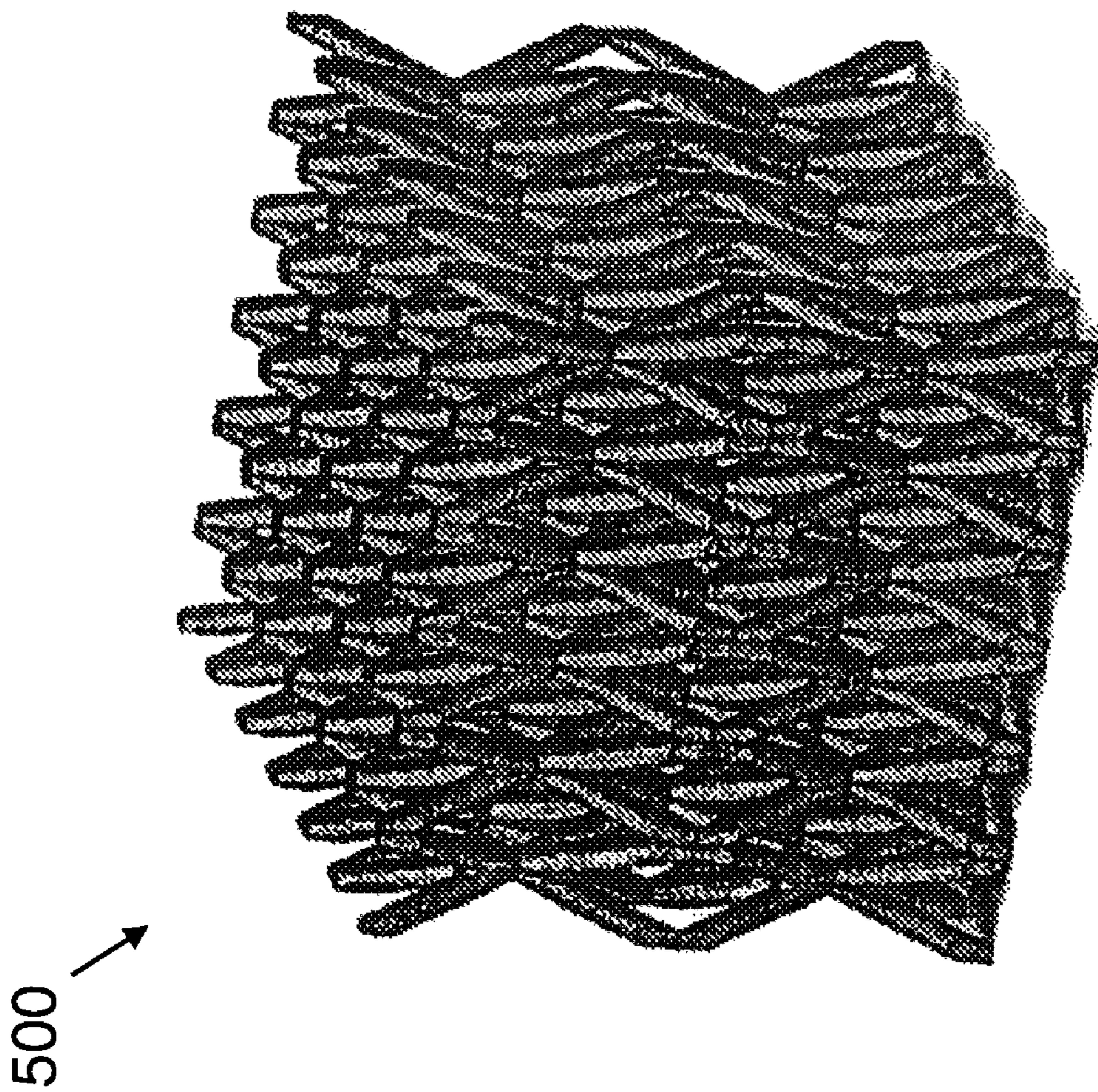


FIG. 5

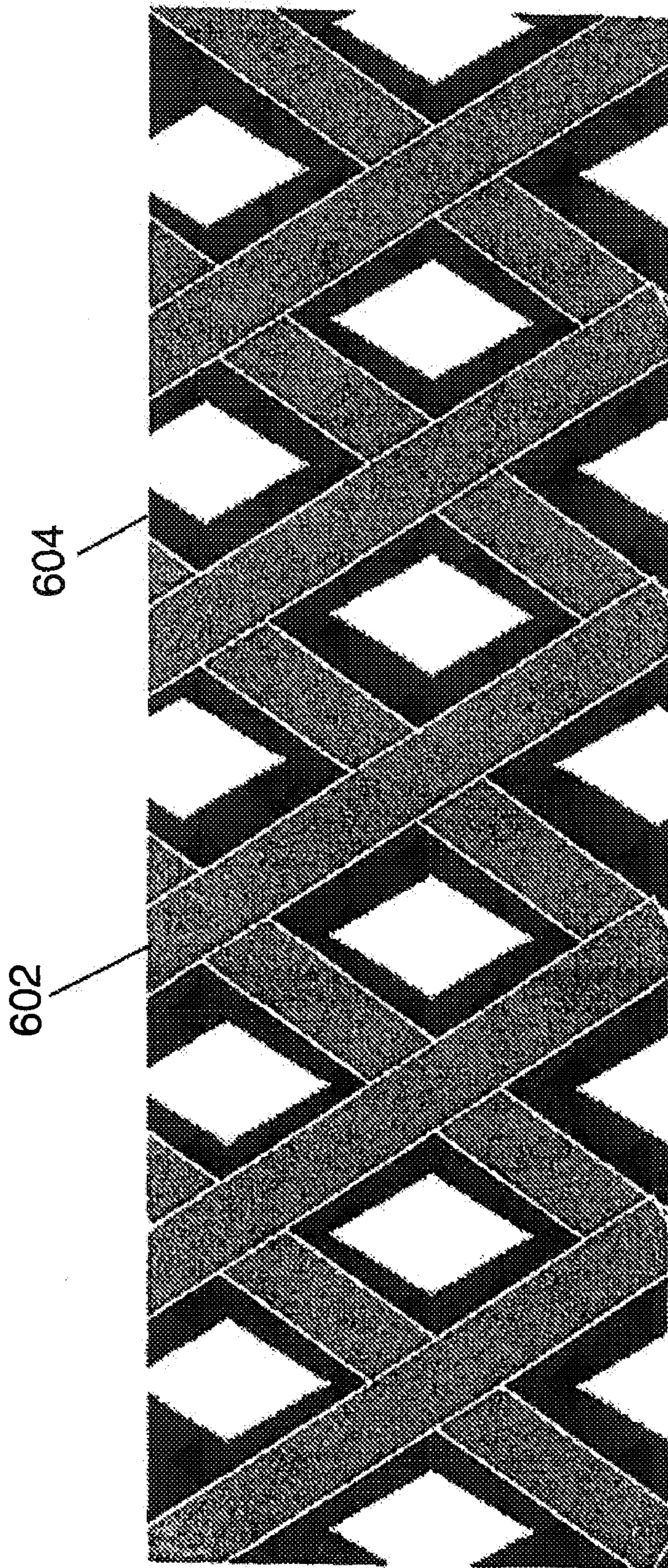


FIG. 6

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**THREE-DIMENSIONAL BIOLOGICAL
SCAFFOLD COMPROMISING POLYMER
WAVEGUIDES**

CROSS-REFERENCE TO RELATED
APPLICATION

This application relates to U.S. patent application Ser. No. 11/580,335, filed on Oct. 13, 2006, now U.S. Pat. No. 7,382,959, entitled "Optically Oriented Three-Dimensional Polymer Microstructures," the entire content of which is incorporated herein by reference.

BACKGROUND OF THE INVENTION

The present invention relates to micro-structures. In particular, the present invention relates to biologically compatible micro-structures and method of making the same.

Scaffold-based biological tissue engineering requires the formation of new tissues which is strongly dependent on the three-dimensional (3D) environment provided by the scaffold. Characteristics of the scaffold that can influence the 3D environment includes its composition, its porous architecture, and its biological response to surrounding tissues/cellular media.

U.S. Pat. No. 6,379,962 (hereinafter the '962 patent) discloses a polymer scaffold having an extensively interconnected macroporous network with macropores having microporous struts as walls. The polymer may be a bio-compatible or bio-degradable polymer. The polymer scaffold is prepared by mixing a liquid polymer with particles, precipitating the liquid polymer with a non-solvent for the liquid polymer and dissolving the particles with a solvent to form the macroporous polymer scaffold. The surface of the polymer scaffold may be modified by acid or base treatment, or by collagen or calcium phosphate deposition. However, this polymer as disclosed in the '962 patent requires surface modification with acid or base treatment or by collagen or calcium phosphate deposition. Furthermore, the structure of the polymer is foam-like with disordered pores that are not homogeneous.

U.S. Pat. No. 6,875,442 (hereinafter the '442 patent) discloses a polymer scaffold with an interconnected macroporous network. The polymer scaffold disclosed embodies macropores having a diameter in a range of 0.5-3.5 mm. The polymer scaffold is prepared using a process which combines the techniques of particulate leaching and phase inversion to render a process that provides amplified means by which to control the morphology of the resulting polymer scaffold. However, the structure disclosed by the '442 patent is foam-like with disordered pores that are not homogeneous.

Similar to the '442 patent, U.S. Pat. No. 7,022,522 (hereinafter the '522 patent) discloses a polymer scaffold that includes an interconnected macroporous network. However, the polymer as disclosed in the '522 patent requires multiple steps to manufacture including particulate leaching and phase inversion. Furthermore, the structure is foam-like with disordered pores that are not homogeneous.

U.S. Pat. No. 6,993,406 (hereinafter the '406 patent) discloses a method for forming a three-dimensional biocompatible porous scaffold structure using a solid freeform fabrication technique that can be used as a medical implant into a living organism. Imaging technology and analysis is first used to determine the three-dimensional design required for the medical implant, such as a bone implant or graft, fashioned as a three-dimensional, biocompatible scaffold structure. The technique is used to either directly produce the three-dimen-

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sional porous scaffold structure or to produce an over-sized three-dimensional porous scaffold lattice which can be machined to produce the designed three-dimensional porous scaffold structure for implantation. The method disclosed by the '406 patent, however, requires pre-fabrication software and can manufacture bio-compatible ceramics only.

Therefore, it is desirable to provide an ordered 3D biological scaffold that can be manufactured in simple steps with ordered interconnected pores with controlled pore size. Furthermore, it is desirable to have an ordered 3D biological scaffold that can enable specific cell/tissue growth.

SUMMARY OF THE INVENTION

In accordance with the embodiments of the present invention, different embodiments of a three-dimensional biological scaffold and a method of making the same are provided.

In one embodiment according to the present invention, a three-dimensional (3D) biological scaffold is provided. The 3D biological scaffold includes at least three sets of bio-degradable polymer waveguides extending along at least three respective directions. The at least three sets of bio-degradable polymer waveguides interpenetrate each other at a plurality of nodes to form a self-supporting structure having a plurality of ordered interconnected pores. The plurality of ordered interconnected pores are sized to facilitate ingress of at least one of biological cells, vascular tissue, or nutrient media.

The plurality of ordered interconnected pores may be sized to about 100-300 μm for trabecular bone implants. A bio-degradable polymer waveguide of the at least three sets of bio-degradable polymer waveguides may have a cross-sectional diameter about 1 μm to about 1 mm. The plurality of ordered interconnected pores may occupy a free space of not less than about 40% by volume and not greater than about 99% by volume of the self-supporting structure. The at least three sets of bio-degradable polymer waveguides may include a bio-degradable polymer selected from one of starches, starch-similar polymers, starch-synthetic blend polymers, esters, amides, anhydrides, urethanes, ureas, carbonates, or saccharides. Surfaces of the three-dimensional biological scaffold may be treated to facilitate cellular recognition and proliferation with a material that has affinity for one of polymeric, ceramic, or metallic surfaces. The material may include one of a silane, a bi-phasic calcium phosphate, a bone morphogenetic protein, a polysaccharide, a biochemical agent, or a mineral precursor.

In another embodiment, a three-dimensional biological scaffold is provided. The three-dimensional biological scaffold includes at least three sets of polymer waveguides extending along at least three respective directions. Surfaces of the at least three sets of polymer waveguides are coated with one or more coating layers. The at least three sets of polymer waveguides interpenetrate each other at a plurality of nodes to form a self-supporting structure having a plurality of ordered interconnected pores. The plurality of ordered interconnected pores are sized to facilitate ingress of at least one of biological cells, vascular tissue, or nutrient media.

The plurality of ordered interconnected pores may be sized to about 250 μm . The one or more coating layers may include one of metal or ceramic. The one or more coating layers may include a material selected from one of titanium (Ti), nickel (Ni), Ti6Al4V, or nickel alloy. The one or more coating layers may include a material selected from one of silicon carbide (SiC), silicon nitride (Si₃N₄), hafnium carbide (HfC), chromium carbide (Cr₃C₂), boron nitride (B₄N), 1:5 cubic boron nitride (c-BN), hexagonal boron nitride (h-BN), boron car-

bide (B4C), aluminum oxide (Al₂O₃), titanium diboride (TiB₂), titanium nitride (TiN), or zirconium dioxide (ZrO₂). A polymer waveguide of the at least three sets of polymer waveguides may have a cross-sectional diameter from about 1 μm to about 1 mm. The polymer waveguide may have a cross-sectional diameter from about 100 μm to about 500 μm. The plurality of ordered interconnected pores may occupy a free space of not less than about 40% by volume and not greater than about 99% by volume of the self-supporting structure. The free space may be not less than about 70% by volume and not greater than about 90% by volume of the self-supporting structure. Surfaces of the three-dimensional biological scaffold may be treated to facilitate cellular recognition and proliferation with a material that has affinity for one of polymeric, ceramic, or metallic surfaces. The material may include one of a silane, a bi-phasic calcium phosphate, a bone morphogenetic protein, a polysaccharide, a biochemical agent, or a mineral precursor.

In still another embodiment, a method for forming a three-dimensional biological scaffold is provided. A volume of photo-monomer is secured. A mask having a plurality of apertures is placed between at least one collimated light source and the volume of photo-monomer. A plurality of collimated light beams from the at least one collimated light source is directed through the plurality of apertures into a portion of the volume of photo-monomer to polymerize the portion to form a plurality of bio-degradable polymer waveguides. And, photo-monomer of the volume of photo-monomer not polymerized is removed to leave behind the three-dimensional biological scaffold having a plurality of ordered interconnected pores. At least three of the plurality of bio-degradable polymer waveguides intersect each other at an aperture of the plurality of apertures. The quantity and opening sizes of the plurality of apertures are selected to adjust the quantity and dimensions of the plurality of ordered interconnected pores to facilitate ingress of at least one of biological cells, vascular tissue, or nutrient media. The plurality of ordered interconnected pores may occupy a free space of not less than about 40% by volume and not greater than about 99% by volume of the three-dimensional biological scaffold.

In yet another embodiment, a method for forming a three-dimensional biological scaffold is provided. A volume of photo-monomer is secured. A mask having a plurality of apertures is placed between at least one collimated light source and the volume of photo-monomer. A plurality of collimated light beams from the at least one collimated light source is directed through the plurality of apertures into a portion of the volume of photo-monomer to polymerize the portion to form a plurality of polymer waveguides. Photo-monomer of the volume of photo-monomer not polymerized is removed to leave behind the three-dimensional biological scaffold having a plurality of ordered interconnected pores. And, surfaces of the plurality of polymer waveguides are covered with one or more coating layers. At least three of the plurality of polymer waveguides intersect each other at an aperture of the plurality of apertures. The quantity and opening sizes of the plurality of apertures are selected to adjust the quantity and dimensions of the plurality of ordered interconnected pores to facilitate ingress of at least one of biological cells, vascular tissue, or nutrient media.

The plurality of ordered interconnected pores may occupy a free space of not less than about 40% by volume and not greater than about 99% by volume of the three-dimensional biological scaffold. The one or more coating layers may include metal, and the one or more coating layers may include ceramic.

BRIEF DESCRIPTION OF THE DRAWINGS

FIG. 1 is a schematic diagram of a system to form a single waveguide from a single collimated beam through a single aperture pursuant to aspects of the present invention.

FIG. 2 is a schematic diagram of a system to form multiple waveguides from a single collimated beam or multiple collimated beams through a single aperture pursuant to aspects of the present invention.

FIG. 3 is a schematic diagram of a system to form a 3D structure (e.g., a 3D ordered polymer microstructure) formed from multiple waveguides created using a single collimated beam or multiple collimated beams through multiple apertures pursuant to aspects of the present invention.

FIG. 4a illustrates an example of a square mask pattern (or a square mask aperture pattern) pursuant to aspects of the present invention.

FIG. 4b illustrates an example of a hexagonal mask pattern (or a hexagonal mask aperture pattern) pursuant to aspects of the present invention.

FIG. 5 illustrates an example of a 3D polymer scaffold pursuant to aspects of the present invention.

FIG. 6 illustrates a cross-sectional view of another example of a 3D polymer scaffold pursuant to aspects of the present invention.

DETAILED DESCRIPTION

The present invention will now be described more fully hereinafter with reference to the accompanying drawings, in which exemplary embodiments of the invention are shown. The dimensions of layers and other elements shown in the accompanying drawings may be exaggerated to more clearly show details. The present invention should not be construed as being limited to the dimensional relations shown in the drawings, nor should the individual elements shown in the drawings be construed to be limited to the dimensions shown.

Embodiments of the present invention provide a 3D biological scaffold, defined by a microtruss structure made of a polymeric, ceramic or metallic material, that has application in biological systems. The 3D biological scaffold may be made solely of a polymer (e.g., bio-degradable or other), ceramic, metal, or any combination thereof with the microtruss structure. An optically templated polymer scaffold determines the final shape and dimensions of the 3D biological scaffold which has the proper pore size and surface chemistry to provides an optimal environment for vascular, cellular, and biomineral growth/remodeling.

An ideal 3D biological scaffold should have appropriate mechanical properties, adequate degradation rate, porosity, interconnectivity, and permeability to allow ingress of cells, vascular tissue, and other nutrient media. The diameter of the pores of the 3D biological scaffold should be at least 100 micron and not larger than 300 micron to enable cell penetration into the structure, to allow tissue ingrowths and to allow blood supply and nutrient to get into the scaffold. For example, the pore size of the 3D biological scaffold may be sized to 250 micron to facilitate trabecular bone implant application. In addition, having the proper surface chemistry is required to enhance cell recognition, attachment, and proliferation. A simple polymerization scheme allows for a controllable, reproducible polymeric structure having a 3D porous network with controlled porosity and surface chemistry. Unlike other known scaffolds in the art, which lack an interconnected porous network, embodiments of the present invention provide an ordered 3D structure with optimized

characteristics, for example, pore scale and interconnectivity, for a particular biological system based on cell diameter, chemistry and other factors.

The present invention has applications in many areas. For example, the medical, dental, and orthopedic communities need biologically compatible structures that provide ideal templating, porosity and interconnectivity for cell, bone, and tooth growth. Embodiments of the present invention can enable polymeric, ceramic, and metallic structures that are ideal or highly compatible for tissue engineering.

Embodiments of the present invention are based on a 3D polymer scaffold. The techniques of making the 3D polymer scaffold are disclosed in a related U.S. patent application Ser. No. 11/580,335 (hereinafter the '335 application) entitled "Optically Oriented Three-Dimensional Polymer Microstructures," which is incorporated herein by reference in its entirety. The '335 application discloses a method and system of fabricating porous cellular polymer materials with an ordered 3D microstructure using a simple technique. These cellular materials are created from a pattern of self-propagating polymer waveguides, which are formed in an appropriate photopolymer. More detail of the techniques can be found in the '335 application.

In reference to the '335 application, a fixed light input (e.g., collimated UV light) is used to cure (i.e., polymerize) polymer optical waveguides, which can self-propagate in a 3D pattern. The propagated polymer optical waveguides form an ordered 3D microstructure that can be polymerized without anything moving during the formation process to provide a path to large scale, inexpensive production.

Some liquid polymers, referred to as photopolymers, undergo a refractive index change during the polymerization process. The refractive index change can lead to a formation of polymer optical waveguides. If a monomer that is photosensitive is exposed to light (e.g., typically UV) under the right conditions, the initial area of polymerization, such as a small circular area, will "trap" the light and guide it to the tip of the polymerized region, further advancing that polymerized region. This process will continue, leading to the formation of a waveguide structure with approximately the same cross-sectional dimensions along its entire length.

The '335 application discloses a polymer cellular material with an ordered 3D microstructure by creating a pattern of self-propagating optical waveguides in an appropriate photopolymer. A formation of a single and multiple polymer waveguides will be described in more detail below, followed by a more detailed description on how to pattern these polymer waveguides to form an ordered 3D microstructure.

Formation of a Single Polymer Waveguide with Single Aperture

Referring to FIG. 1, a system to form a single optical waveguide includes a collimated light source **100**, a reservoir **110** (e.g., a mold) having a volume of monomer **120** that will polymerize at a wavelength of a collimated light beam **170** provided by the light source **100**, and a patterning apparatus, such as a mask **130** with a single aperture **140** (e.g., an open area) of a given shape and dimension.

The single collimated beam **170** is directed through the aperture **140** in the mask **130** to the monomer **120**. Between the mask **130** and the monomer **120**, there may be a substrate **150**. The substrate can be composed of a material, such as glass, Mylar, and other suitable materials that will transmit the incident light beam to the monomer **120**. That is, in one embodiment of the present invention, the substrate **150** is substantially transparent to the incident light beam. On the

surface of the monomer **120**, in the area exposed to a portion of the light beam, an optical waveguide **160** will begin to polymerize.

The index of refraction change between the polymer and monomer will "trap" and "focus" the light in the polymer and guide the polymerization process. Due to this self-guiding/self-focusing effect, the polymerized waveguide **160** will form with an approximately constant cross-section and a length much greater than the cross-sectional dimensions. The direction in which this polymer waveguide **160** will grow is dependent on the direction of the incident beam. The cross-section of the polymer waveguide **160** is dependent on the shape and dimensions of the incident collimated beam, which in turn is dependent on the shape and dimensions of the aperture **140** in the mask **130**. The length to which the polymer waveguide **160** can "grow" is dependent on a number of parameters including the size, intensity, and exposure time of the incident beam, as well as the light absorption/transmission properties of the photopolymer. The time in which it takes to form a polymer waveguide depends on the kinetics of the polymerization process.

To put it another way, when the portion of the collimated light beam passes through the mask **130** and first hits the liquid photo-monomer **120**, a polymer "tip" is formed. There is a large enough difference between the refractive index of the monomer and the polymer to cause internal reflection of the light in the polymer—this is the same principle as when light travels through fiber optics. Because of this internal reflection effect, the light is essentially focused to the tip of the polymer, causing the monomer at the tip to cure (i.e., polymerize). This will also propagate the tip of the polymer through the liquid monomer **120**, forming the self-propagating polymer optical waveguide **160**. In addition, because of this internal reflection effect, the waveguide **160** can be "very" long with respect to the cross-sectional dimensions, all while maintaining a constant cross-section through its length. Eventually the formation of the polymer waveguide **160** will stop at the end of the monomer reservoir **110**, or it will stop prior to that if there is not enough energy to polymerize the monomer **120**. This happens because the polymer itself will absorb some of the portion of the collimated light beam passing through the mask **130**.

Formation of Multiple Polymer Waveguides with Single Aperture

As mentioned above, the direction in which the polymer waveguide will form is dependent on the angle of the incident collimated beam. If the collimated beam is perpendicular to a flat monomer surface (as shown in FIG. 1), the polymer waveguide will propagate, or grow perpendicular to the monomer surface. By contrast, referring to FIG. 2, if the incident collimated beam is directed at an angle, the polymer waveguide will grow at an angle relative to the monomer surface. Note this angle will be affected by the change in refractive index between the air and/or substrate and the monomer due to refraction.

That is, as shown in FIG. 2, a system to form multiple optical waveguides **260** includes one or more collimated light sources **200**, a reservoir **210** (e.g., mold) having a volume of monomer **220** that will polymerize at a wavelength of collimated light beams provided by the light sources **200**, and a patterning apparatus, such as a mask **230** with a single aperture (e.g., open area) **240** of a given shape and dimension. Between the mask **230** and the monomer **220**, there may be a substrate **250**.

Through the single aperture **240** as described above, the multiple waveguides **260** can be formed by directing multiple collimated beams at different angles through the aperture

240. That is, in one embodiment of the invention, a single collimated light source is used. Multiple waveguides are formed from a plurality of exposures of the collimated light beam of the single collimated light source with a mask having a single aperture, and the single collimated light source is adapted to move with respect to the mask between each of the exposures.

Alternatively, the multiple waveguides **260** can be formed one at a time through the single aperture **240** by simply changing the incident angle of a single collimated beam after the formation of each of the waveguides **240**. That is, in another embodiment of the present invention, multiple collimated light sources are adapted to produce and direct multiple collimated light beams at different angles with respect to a point of a mask having a single aperture. Multiple waveguides are formed from a single exposure of the multiple light beams of the multiple collimated light sources with the mask having the single aperture.

Formation of 3D Microstructure Using Patterned Optical Waveguides

The technique to create a 3D polymer microstructure is based on the above described approach for forming multiple optical waveguides with a single aperture. However, instead of using a mask with a single aperture, a mask with a two-dimensional pattern of apertures is used to create a three-dimensional polymer microstructure as is shown in FIG. **3**.

Referring to FIG. **3**, a system to form a 3D polymer microstructure includes one or more collimated light sources **300**, a reservoir (e.g., a mold) **310** having a volume of monomer **320** that will polymerize at a wavelength of collimated light beams provided by the light sources **300**, and a patterning apparatus, such as a mask **330** with multiple apertures (e.g., open areas) **340**. Each of the apertures **340** has a given shape and dimension substantially matching a cross section geometry of a polymer waveguide **360**. Between the mask **330** and the monomer **320**, there may be a substrate **350**. Here, in FIG. **3**, a truly 3D network can be formed because the intersecting polymer waveguides **360** will simply polymerize together, but will not interfere with waveguide propagation. Also, the spacing between the plurality of waveguides **360** corresponds with the pattern of the plurality of apertures **340**. The pattern of the apertures **340** may, for example, be in a square pattern as shown in FIG. **4a** and/or in a hexagonal pattern as shown in FIG. **4b**. The hole (i.e., aperture) spacing, i.e., distance between apertures **340** in the mask **330**, and the number of waveguides **360** formed from each of the apertures **340** will determine the open volume fraction (i.e., open space) of the formed 3D microstructure. It should be appreciated by a person skilled in the art that once a certain pore size is determined for a particular application, the hole spacing and the size of the aperture can be adjusted accordingly to produce the desired pore size.

As such, through the process shown in FIG. **3**, an exemplary 3D polymer scaffold **500** (e.g., a 3D ordered polymer microstructure) can be designed and provided as shown in FIG. **5**. The design parameters include: 1) the angle and pattern of the polymer waveguides with respect to one another; 2) the packing, or relative density of the resulting cellular structure (or the open volume fraction); and 3) the cross-sectional shape and dimensions of the polymer waveguides. By varying these design parameters, the exemplary 3D polymer scaffold **500** can be adapted to facilitate a particular implant application. By way of example, in human trabecular bone implant application, the pore size of the scaffold can be within the range of 0.5 to 3.5 mm to match the pore size of human trabecular bone. The high level of intercon-

nectivity of the pores of the 3D polymer scaffold **500** enhances both penetration of the scaffold **500** by cells and nutrient flow to the cells.

In one embodiment according to the present invention, the 3D polymer scaffold **500** includes bio-degradable polymer such as, but not limited to, starches, starch-similar polymers, starch-synthetic blend polymers, esters, amides, anhydrides, urethanes, ureas, carbonates, or saccharides.

In another embodiment according to the present invention, the truss elements (i.e., the waveguides **360**) may have diameters between about 1 μm and about 1 mm, and the 3D polymer scaffold can have between 40% and 99% of interconnected free space.

In still another embodiment according to the present invention, the 3D polymer scaffold **602** can be deposited with one or more layers of material **604** as shown in FIG. **6**. The layers of material can be, but not limited to, ceramic or metal. Exemplary ceramic can be, but not limited to, one of silicon carbide (SiC), silicon nitride (Si₃N₄), hafnium carbide (HfC), chromium carbide (Cr₃C₂), boron nitride (B₄N), cubic boron nitride (c-BN), hexagonal boron nitride (h-BN), boron carbide (B₄C), aluminum oxide (Al₂O₃), titanium diboride (TiB₂), titanium nitride (TiN), or zirconium dioxide (ZrO₂). Exemplary metal can be, but not limited to, one of titanium (Ti), nickel (Ni), Ti₆Al₄V, or nickel alloy. The layers of material can be applied by chemical vapor deposition, reacting the 3D polymer scaffold with a gas or liquid phase reactant, or coating the 3D polymer scaffold with a suitable pre-ceramic polymer.

In yet another embodiment according to the present invention, the 3D polymer scaffold's surface is treated to enable specific functionalities, such as specific cellular recognition and proliferation needed to stimulate cell growth and achieve bio-compatibility. By immersing a ceramic or metal coated 3D polymer scaffold in a solution bath of various mono- or multi-functional molecules (e.g., biochemical agents or mineral precursor), a biologically specific surface can be produced. The functional molecules can have an affinity for polymeric, ceramic, or metallic surfaces. This treated 3D polymer scaffold can provide bone-implant scaffolds absorption/recognition sites for molecular/cellular species such as bone morphogenetic proteins and bi-phasic calcium phosphate that are critical to osteogenesis. These molecules/minerals are critical during osteoinduction (i.e., allows for cell differentiation and bone morphogenesis). In addition, pore size and surface functionality are critical during osteogenesis (i.e., forming bone), whereby proper adsorption sites and transportation pathways are provided by the 3D scaffold to the bone-mineralizing cells (i.e., osteoblasts). Cell transport is also critical during remodeling of bone; that is, it is necessary to transport bone-resorbing cells (i.e., osteoclasts) to growing/remodeled bone.

Although certain exemplary embodiments of the present invention have been disclosed for illustrative purposes, those skilled in the art would appreciate that various modifications, additions and subtractions are possible, without departing from the scope and spirit of the present invention as disclosed in the accompanying claims and their equivalents.

What is claimed is:

1. A three-dimensional biological scaffold comprising:
 - at least three sets of bio-degradable polymer waveguides extending along at least three different directions,
 - wherein the at least three sets of bio-degradable polymer waveguides interpenetrate each other at a plurality of nodes to form a self-supporting structure having a plurality of ordered interconnected pores, at least three of

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- the bio-degradable polymer waveguides intersecting each other at a node of the plurality of nodes at non-perpendicular angles, and
 wherein the plurality of ordered interconnected pores are sized to facilitate ingress of at least one of biological cells, vascular tissue, or nutrient media.
2. The three-dimensional biological scaffold of claim 1, wherein the plurality of ordered interconnected pores are sized to about 100 μm to 300 μm .
3. The three-dimensional biological scaffold of claim 1, wherein a bio-degradable polymer waveguide of the at least three sets of bio-degradable polymer waveguides has a cross-sectional diameter about 1 μm to about 1 mm.
4. The three-dimensional biological scaffold of claim 1, wherein the plurality of ordered interconnected pores occupy a free space of not less than about 40% by volume and not greater than about 99% by volume of the self-supporting structure.
5. The three-dimensional biological scaffold of claim 1, wherein the at least three sets of bio-degradable polymer waveguides comprise a bio-degradable polymer selected from one of starches, starch-similar polymers, starch-synthetic blend polymers, esters, amides, anhydrides, urethanes, ureas, carbonates, or saccharides.
6. The three-dimensional biological scaffold of claim 1, wherein surfaces of the three-dimensional biological scaffold are treated with a material selected from the group consisting of a silane, a biphasic calcium phosphate, a bone morphogenetic protein, and a polysaccharide.
7. A three-dimensional biological scaffold comprising:
 at least three sets of polymer waveguides extending along at least three different directions; and
 one or more coating layers on surfaces of the at least three sets of polymer waveguides,
 wherein the at least three sets of polymer waveguides interpenetrate each other at a plurality of nodes to form a self-supporting structure having a plurality of ordered interconnected pores, at least three of the polymer waveguides intersecting each other at a node of the plurality of nodes at non-perpendicular angles, and
 wherein the plurality of ordered interconnected pores are sized to facilitate ingress of at least one of biological cells, vascular tissue, or nutrient media.

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8. The three-dimensional biological scaffold of claim 7, wherein the plurality of ordered interconnected pores are sized to about 100 μm to 300 μm .
9. The three-dimensional biological scaffold of claim 7, wherein the one or more coating layers comprise one of metal or ceramic.
10. The three-dimensional biological scaffold of claim 9, wherein the one or more coating layers comprise a material selected from one of titanium (Ti), nickel (Ni), Ti6Al4V, or nickel alloy.
11. The three-dimensional biological scaffold of claim 9, wherein the one or more coating layers comprise a material selected from one of silicon carbide (SiC), silicon nitride (Si₃N₄), hafnium carbide (HfC), chromium carbide (Cr₃C₂), boron nitride (B₄N), cubic boron nitride (c-BN), hexagonal boron nitride (h-BN), boron carbide (B₄C), aluminum oxide (Al₂O₃), titanium diboride (TiB₂), titanium nitride (TiN), or zirconium dioxide (ZrO₂).
12. The three-dimensional biological scaffold of claim 7, wherein a polymer waveguide of the at least three sets of polymer waveguides has a cross-sectional diameter from about 1 μm to about 1 mm.
13. The three-dimensional biological scaffold of claim 12, wherein the polymer waveguide has a cross-sectional diameter from about 100 μm to about 500 μm .
14. The three-dimensional biological scaffold of claim 7, wherein the plurality of ordered interconnected pores occupy a free space of not less than about 40% by volume and not greater than about 99% by volume of the self-supporting structure.
15. The three-dimensional biological scaffold of claim 14, wherein the free space is not less than about 70% by volume and not greater than about 90% by volume of the self-supporting structure.
16. The three-dimensional biological scaffold of claim 7, wherein surfaces of the three-dimensional biological scaffold are treated with a material selected from the group consisting of a silane, a bi-phasic calcium phosphate, a bone morphogenetic protein, and a polysaccharide.

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